Welcome to STN International! Enter x:x

LOGINID: SSSPTA1642BJF

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

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* * * * * * * * * *
                     Welcome to STN International
                 Web Page for STN Seminar Schedule - N. America
NEWS
         JAN 02
NEWS
                 STN pricing information for 2008 now available
NEWS
         JAN 16
                 CAS patent coverage enhanced to include exemplified
                 prophetic substances
NEWS
         JAN 28
                 USPATFULL, USPAT2, and USPATOLD enhanced with new
                 custom IPC display formats
NEWS
         JAN 28
                 MARPAT searching enhanced
NEWS
         JAN 28
                 USGENE now provides USPTO sequence data within 3 days
                 of publication
NEWS
         JAN 28
                 TOXCENTER enhanced with reloaded MEDLINE segment
NEWS 8
         JAN 28
                 MEDLINE and LMEDLINE reloaded with enhancements
NEWS 9
         FEB 08
                 STN Express, Version 8.3, now available
NEWS 10 FEB 20
                 PCI now available as a replacement to DPCI
NEWS 11 FEB 25
                 IFIREF reloaded with enhancements
NEWS 12 FEB 25
                 IMSPRODUCT reloaded with enhancements
NEWS 13 FEB 29
                 WPINDEX/WPIDS/WPIX enhanced with ECLA and current
                 U.S. National Patent Classification
                 IFICDB, IFIPAT, and IFIUDB enhanced with new custom
NEWS 14
         MAR 31
                 IPC display formats
NEWS 15
         MAR 31
                 CAS REGISTRY enhanced with additional experimental
NEWS 16
                 CA/CAplus and CASREACT patent number format for U.S.
         MAR 31
                 applications updated
NEWS 17
         MAR 31
                 LPCI now available as a replacement to LDPCI
NEWS 18
         MAR 31
                 EMBASE, EMBAL, and LEMBASE reloaded with enhancements
NEWS 19
         APR 04
                 STN AnaVist, Version 1, to be discontinued
NEWS 20 APR 15
                 WPIDS, WPINDEX, and WPIX enhanced with new
                 predefined hit display formats
NEWS 21 APR 28
                 EMBASE Controlled Term thesaurus enhanced
NEWS 22 APR 28
                 IMSRESEARCH reloaded with enhancements
NEWS 23 MAY 30
                 INPAFAMDB now available on STN for patent family
                 searching
NEWS 24
         MAY 30
                 DGENE, PCTGEN, and USGENE enhanced with new homology
                 sequence search option
NEWS 25
         JUN 06
                 EPFULL enhanced with 260,000 English abstracts
NEWS 26
         JUN 06
                 KOREAPAT updated with 41,000 documents
NEWS 27
         JUN 13
                 USPATFULL and USPAT2 updated with 11-character
                 patent numbers for U.S. applications
NEWS 28
         JUN 19
                 CAS REGISTRY includes selected substances from
                 web-based collections
NEWS 29
         JUN 25
                 CA/CAplus and USPAT databases updated with IPC
                 reclassification data
NEWS 30
         JUN 30
                 AEROSPACE enhanced with more than 1 million U.S.
                 patent records
NEWS 31
         JUN 30
                 EMBASE, EMBAL, and LEMBASE updated with additional
                 options to display authors and affiliated
```

organizations

NEWS 32 JUN 30 STN on the Web enhanced with new STN AnaVist Assistant and BLAST plug-in

NEWS 33 JUN 30 STN AnaVist enhanced with database content from EPFULL

NEWS EXPRESS JUNE 27 08 CURRENT WINDOWS VERSION IS V8.3, AND CURRENT DISCOVER FILE IS DATED 23 JUNE 2008.

NEWS HOURS STN Operating Hours Plus Help Desk Availability

NEWS LOGIN Welcome Banner and News Items

NEWS IPC8 For general information regarding STN implementation of IPC 8

Enter NEWS followed by the item number or name to see news on that specific topic.

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FILE 'HOME' ENTERED AT 08:59:51 ON 07 JUL 2008

=> file reg COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.21 0.21

FULL ESTIMATED COST

FILE 'REGISTRY' ENTERED AT 09:00:16 ON 07 JUL 2008
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
COPYRIGHT (C) 2008 American Chemical Society (ACS)

Property values tagged with IC are from the ${\tt ZIC/VINITI}$ data file provided by InfoChem.

STRUCTURE FILE UPDATES: 6 JUL 2008 HIGHEST RN 1032827-24-9 DICTIONARY FILE UPDATES: 6 JUL 2008 HIGHEST RN 1032827-24-9

New CAS Information Use Policies, enter HELP USAGETERMS for details.

TSCA INFORMATION NOW CURRENT THROUGH January 9, 2008.

Please note that search-term pricing does apply when conducting SmartSELECT searches.

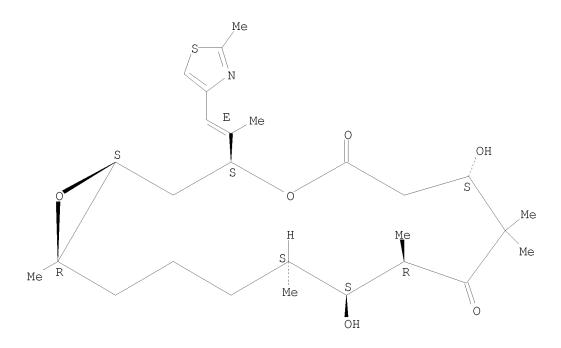
REGISTRY includes numerically searchable data for experimental and predicted properties as well as tags indicating availability of experimental property data in the original document. For information on property searching in REGISTRY, refer to:

http://www.cas.org/support/stngen/stndoc/properties.html

```
=> E "EPOTHILONE B"/CN 25
E1
            1 EPOTHILONE A8/CN
E2
            1
                 EPOTHILONE A9/CN
E3
            1 --> EPOTHILONE B/CN
E4
           1 EPOTHILONE B (12R, 13R) ACETONIDE/CN
                EPOTHILONE B A-EPOXIDE/CN
E.5
           1
Ε6
            1
                EPOTHILONE B ACID/CN
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EPOTHILONE B HYDROXYLASE/CN
EPOTHILONE B HYDROXYLASE (AMYCOLATOPSIS ORIENTALIS GENE EBH)/CN
EPOTHILONE B N-OXIDE/CN
EPOTHILONE B10/CN
EPOTHILONE C/CN
EPOTHILONE C/CN
EPOTHILONE C BIS(TERT-BUTYLDIMETHYLSILYL) ETHER/CN
EPOTHILONE C/D 12,13-EPOXIDASE/CN
EPOTHILONE C/D MONOOXYGENASE/CN
EPOTHILONE C/D SYNTHETASE/CN
EPOTHILONE C1/CN
EPOTHILONE C2/CN
EPOTHILONE C3/CN
EPOTHILONE C3/CN
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EPOTHILONE C9/CN
EPOTHILONE C9/CN
EPOTHILONE C9/CN
EPOTHILONE D/CN
E7
E.8
E9
E10
E11
E12
E13
E14
E15
E16
E17
E18
E19
E20
E21
E22
E23
E24
E25
=> S E3
L1
                      1 "EPOTHILONE B"/CN
=> S L1 EXA SAM
SAMPLE IS IGNORED AS A SCOPE FOR THIS SEARCH
                      1 "EPOTHILONE B"/CN
=> DIS L2 1 SAM
THE ESTIMATED COST FOR THIS REQUEST IS 1.04 U.S. DOLLARS
DO YOU WANT TO CONTINUE WITH THIS REQUEST? (Y)/N:Y
        ANSWER 1 OF 1 REGISTRY COPYRIGHT 2008 ACS on STN
L2
         4,17-Dioxabicyclo[14.1.0]heptadecane-5,9-dione, 7,11-dihydroxy-
ΙN
         8, 8, 10, 12, 16-pentamethyl-3-[(1E)-1-methyl-2-(2-methyl-4-thiazolyl)ethenyl]-
         , (1S,3S,7S,10R,11S,12S,16R)-
        C27 H41 N O6 S
MF
Absolute stereochemistry. Rotation (-).
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Double bond geometry as shown.



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

=>

---Logging off of STN---

=>

Executing the logoff script...

=> LOG Y

COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 13.64 13.85

FULL ESTIMATED COST

STN INTERNATIONAL LOGOFF AT 09:03:08 ON 07 JUL 2008

Connecting via Winsock to STN

Welcome to STN International! Enter x:x

LOGINID:SSSPTA1642BJF

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

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Welcome to STN International
                 Web Page for STN Seminar Schedule - N. America
NEWS
NEWS
         JAN 02
                 STN pricing information for 2008 now available
NEWS
        JAN 16
                 CAS patent coverage enhanced to include exemplified
                 prophetic substances
        JAN 28
NEWS
                 USPATFULL, USPAT2, and USPATOLD enhanced with new
                 custom IPC display formats
        JAN 28
                 MARPAT searching enhanced
NEWS
        JAN 28
NEWS
                 USGENE now provides USPTO sequence data within 3 days
                 of publication
NEWS
         JAN 28
                 TOXCENTER enhanced with reloaded MEDLINE segment
NEWS 8
        JAN 28
                 MEDLINE and LMEDLINE reloaded with enhancements
NEWS
        FEB 08
                 STN Express, Version 8.3, now available
NEWS 10
        FEB 20
                 PCI now available as a replacement to DPCI
NEWS 11
        FEB 25
                 IFIREF reloaded with enhancements
NEWS 12
        FEB 25
                 IMSPRODUCT reloaded with enhancements
NEWS 13
        FEB 29
                 WPINDEX/WPIDS/WPIX enhanced with ECLA and current
                 U.S. National Patent Classification
NEWS 14
        MAR 31
                 IFICDB, IFIPAT, and IFIUDB enhanced with new custom
                 IPC display formats
NEWS 15
        MAR 31
                 CAS REGISTRY enhanced with additional experimental
                 spectra
        MAR 31
                 CA/CAplus and CASREACT patent number format for U.S.
NEWS 16
                 applications updated
NEWS 17
        MAR 31
                 LPCI now available as a replacement to LDPCI
NEWS 18
        MAR 31
                 EMBASE, EMBAL, and LEMBASE reloaded with enhancements
NEWS 19
        APR 04
                 STN AnaVist, Version 1, to be discontinued
NEWS 20
        APR 15
                 WPIDS, WPINDEX, and WPIX enhanced with new
                 predefined hit display formats
NEWS 21
        APR 28
                 EMBASE Controlled Term thesaurus enhanced
NEWS 22
        APR 28
                 IMSRESEARCH reloaded with enhancements
NEWS 23
        MAY 30
                 INPAFAMDB now available on STN for patent family
                 searching
NEWS 24
        MAY 30
                 DGENE, PCTGEN, and USGENE enhanced with new homology
                 sequence search option
NEWS 25
         JUN 06
                 EPFULL enhanced with 260,000 English abstracts
                 KOREAPAT updated with 41,000 documents
         JUN 06
NEWS 27
                 USPATFULL and USPAT2 updated with 11-character
         JUN 13
                 patent numbers for U.S. applications
NEWS 28
         JUN 19
                CAS REGISTRY includes selected substances from
                 web-based collections
NEWS 29
         JUN 25
                 CA/CAplus and USPAT databases updated with IPC
                 reclassification data
NEWS 30
         JUN 30
                 AEROSPACE enhanced with more than 1 million U.S.
                 patent records
                 EMBASE, EMBAL, and LEMBASE updated with additional
NEWS 31
         JUN 30
                 options to display authors and affiliated
                 organizations
NEWS 32
         JUN 30
                 STN on the Web enhanced with new STN AnaVist
                 Assistant and BLAST plug-in
NEWS 33
        JUN 30
                STN AnaVist enhanced with database content from EPFULL
NEWS EXPRESS JUNE 27 08 CURRENT WINDOWS VERSION IS V8.3,
             AND CURRENT DISCOVER FILE IS DATED 23 JUNE 2008.
NEWS HOURS
              STN Operating Hours Plus Help Desk Availability
NEWS LOGIN
              Welcome Banner and News Items
NEWS IPC8
              For general information regarding STN implementation of IPC 8
```

Enter NEWS followed by the item number or name to see news on that

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FILE 'HOME' ENTERED AT 10:09:50 ON 07 JUL 2008

=> file pctfull

COST IN U.S. DOLLARS
SINCE FILE TOTAL
ENTRY SESSION
FULL ESTIMATED COST
0.21
0.21

FILE 'PCTFULL' ENTERED AT 10:10:27 ON 07 JUL 2008 COPYRIGHT (C) 2008 Univentio

FILE LAST UPDATED: 4 JUL 2008 <20080704/UP>

FILE COVERS 1978 TO DATE

>>> IMAGES ARE AVAILABLE ONLINE AND FOR EMAIL-PRINTS <<<

>>> NEW FIELD UPTX, FIELD /EW NO LONGER AVAILBLE - SEE HELP CHANGE <<<

=> s epothilon?

L1 2484 EPOTHILON?

=> s 11/ab or 11/ti

144 EPOTHILON?/AB 129 EPOTHILON?/TI

L2 159 (EPOTHILON?/AB) OR (EPOTHILON?/TI)

=> s 12 not py>2001 817323 PY>2001

L3 53 L2 NOT PY>2001

=> s combination and 13

567168 COMBINATION 264042 COMBINATIONS 617900 COMBINATION

(COMBINATION OR COMBINATIONS)

L4 33 COMBINATION AND L3

=> d ibib 1-5

L4 ANSWER 1 OF 33 PCTFULL COPYRIGHT 2008 Univentio on STN

ACCESSION NUMBER: 2001092255 PCTFULL ED 20020826

TITLE (ENGLISH): EPOTHILONE DERIVATIVES AND METHODS FOR MAKING

AND USING THE SAME

TITLE (FRENCH): DERIVES D'EPOTHILONE, PROCEDES DE PRODUCTION

ET METHODES D'UTILISATION

INVENTOR(S): SANTI, Daniel;

FARDIS, Maria; ASHLEY, Gary

PATENT ASSIGNEE(S): KOSAN BIOSCIENCES, INC.;

SANTI, Daniel; FARDIS, Maria; ASHLEY, Gary

DOCUMENT TYPE: Patent

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PATENT INFORMATION:
                       NUMBER
                                KIND DATE
                       ______
                       WO 2001092255 A2 20011206
DESIGNATED STATES
                       AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR
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                       CU CZ DE DK DM DZ EC EE ES FI GB GD GE GH GM HR HU ID
                       IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD
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                       US 2000-60/207,655 20000526
PRIORITY INFO.:
                       US 2000-60/218,260
                                             20000714
                       US 2000-60/231,552 20000911
                       WO 2001-US15763 A 20010515
APPLICATION INFO.:
      ANSWER 2 OF 33
                       PCTFULL COPYRIGHT 2008 Univentio on STN
ACCESSION NUMBER:
                      2001083800 PCTFULL ED 20020826
TITLE (ENGLISH):
                      PRODUCTION OF POLYKETIDES
TITLE (FRENCH):
                      PRODUCTION DE POLYKETIDES
INVENTOR(S):
                       ARSLANIAN, Robert, L.;
                       ASHLEY, Gary;
                       FRYKMAN, Scott;
                       JULIEN, Bryan;
                       KATZ, Leonard;
                       KHOSLA, Chaitan;
                       LAU, Janice;
                       LICARDI, Peter, J.;
                       REGENTIN, Rika;
                       SANTI, Daniel;
                       TANG, Li
                       KOSAN BIOSCIENCES, INC.;
PATENT ASSIGNEE(S):
                       ARSLANIAN, Robert, L.;
                       ASHLEY, Gary;
                       FRYKMAN, Scott;
                       JULIEN, Bryan;
                       KATZ, Leonard;
                       KHOSLA, Chaitan;
                       LAU, Janice;
                       LICARDI, Peter, J.;
                       REGENTIN, Rika;
                       SANTI, Daniel;
                       TANG, Li
DOCUMENT TYPE:
                       Patent
PATENT INFORMATION:
                              KIND DATE
                       NUMBER
                       WO 2001083800 A2 20011108
DESIGNATED STATES
                       AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU
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                       US 2000-09/560,367 20000428
PRIORITY INFO.:
                       US 2000-60/232,696 20000914
US 2000-60/257,517 20001221
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US 2001-09/825,856 20010403
US 2001-09/825,876 20010403
US 2001-60/269,020 20010413
                        WO 2001-US13793 A 20010426
APPLICATION INFO.:
       ANSWER 3 OF 33
                        PCTFULL COPYRIGHT 2008 Univentio on STN
ACCESSION NUMBER:
                        2001081341 PCTFULL ED 20020826
TITLE (ENGLISH):
                        9-OXA-EPOTHILON DERIVATIVES, METHOD FOR THE
                        PRODUCTION AND USE THEREOF IN PHARMACEUTICAL
                        PREPARATIONS
TITLE (FRENCH):
                        DERIVES DE 9-OXA-EPOTHILONE, LEUR PROCEDE DE
                        PRODUCTION ET LEUR UTILISATION PHARMACEUTIQUE
INVENTOR(S):
                        SCHWEDE, Wolfgang;
                        KLAR, Ulrich;
                         SKUBALLA, Werner;
                        BUCHMANN, Bernd;
                        HOFFMANN, Jens;
                        LICHTNER, Rosemarie
                        SCHERING AKTIENGESELLSCHAFT;
PATENT ASSIGNEE(S):
                         SCHWEDE, Wolfgang;
                        KLAR, Ulrich;
                         SKUBALLA, Werner;
                         BUCHMANN, Bernd;
                        HOFFMANN, Jens;
                        LICHTNER, Rosemarie
DOCUMENT TYPE:
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PATENT INFORMATION:
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                                          KIND DATE
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                        WO 2001081341 A2 20011101
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### PRIORITY INFO:: DE 2000-100 20 899.1 20000420 APPLICATION INFO:: WO 2001-EP4551
                        CI CM GA GN GW ML MR NE SN TD TG
       ANSWER 4 OF 33
                        PCTFULL COPYRIGHT 2008 Univentio on STN
ACCESSION NUMBER: 2001073103 PCTFULL ED 20020822 TITLE (ENGLISH): PREPARATION OF EPOTHILONE INTERMEDIATES
TITLE (FRENCH):
                       PREPARATION D'INTERMEDIAIRES D'EPOTHILONE
INVENTOR(S):
                        VITE, Gregory, D.;
                        KIM, Soong-Hoon;
                        HOeEFLE, Gerhard
                        BRISTOL-MYERS SQUIBB COMPANY;
PATENT ASSIGNEE(S):
                        VITE, Gregory, D.;
                        KIM, Soong-Hoon;
                        HOeEFLE, Gerhard
DOCUMENT TYPE:
                        Patent
PATENT INFORMATION:
                                    KIND DATE
                        NUMBER
                        WO 2001073103 A2 20011004
DESIGNATED STATES
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                        CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL
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PRIORITY INFO.:
                       US 2000-60/191,975 20000324
APPLICATION INFO.:
                       WO 2001-US9620
                                           A 20010323
      ANSWER 5 OF 33
                       PCTFULL COPYRIGHT 2008 Univentio on STN
ACCESSION NUMBER:
                       2001070716 PCTFULL ED 20020822
                       A PROCESS FOR THE PREPARATION OF EPOTHILONE
TITLE (ENGLISH):
                       ANALOGS AND INTERMEDIATES
TITLE (FRENCH):
                       PREPARATION D'ANALOGUES ET D'INTERMEDIAIRES D'
                       EPOTHILONE
INVENTOR(S):
                       LI, Wen, Sen;
                       THORNTON, John, E.;
                       GUO, Zhenrong;
                       SWAMINATHAN, Shankar;
                       MCCONLOGUE, Gary, W.
                       BRISTOL-MYERS SQUIBB COMPANY;
PATENT ASSIGNEE(S):
                       LI, Wen, Sen;
                       THORNTON, John, E.;
                       GUO, Zhenrong;
                       SWAMINATHAN, Shankar;
                       MCCONLOGUE, Gary, W.
DOCUMENT TYPE:
                       Patent
PATENT INFORMATION:
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                       WO 2001070716 A1 20010927
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APPLICATION INFO.:
PRIORITY INFO.:
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                       WO 2001-US7749 A 20010312
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     (FILE 'HOME' ENTERED AT 10:09:50 ON 07 JUL 2008)
     FILE 'PCTFULL' ENTERED AT 10:10:27 ON 07 JUL 2008
          2484 S EPOTHILON?
L1
L2
           159 S L1/AB OR L1/TI
L3
            53 S L2 NOT PY>2001
            33 S COMBINATION AND L3
T.4
=> s 14 and (taxol or paclitaxel)
          9622 TAXOL
           272 TAXOLS
          9705 TAXOL
                (TAXOL OR TAXOLS)
         10390 PACLITAXEL
           72 PACLITAXELS
         10392 PACLITAXEL
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(PACLITAXEL OR PACLITAXELS)

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T<sub>1</sub>5
          29 L4 AND (TAXOL OR PACLITAXEL)
=> s 15 and Her?
       988529 HER?
1.6
           29 L5 AND HER?
=> s 15 and (HER2 or HER-2)
         4722 HER2
        118696 HER
         1043 HERS
       119313 HER
                (HER OR HERS)
      1276185 2
         3260 HER-2
               (HER(W)2)
L7
            1 L5 AND (HER2 OR HER-2)
=> d ibib abs
      ANSWER 1 OF 1
                       PCTFULL
                                COPYRIGHT 2008 Univentio on STN
                     1999002514 PCTFULL ED 20020515
ACCESSION NUMBER:
TITLE (ENGLISH):
                       EPOTHILONE DERIVATIVES
TITLE (FRENCH):
                       DERIVES D'EPOTHILONE
INVENTOR(S):
                       VITE, Gregory, D.;
                       BORZILLERI, Robert, M.;
                       KIM, Soong-Hoon;
                       JOHNSON, James, A.
PATENT ASSIGNEE(S):
                       BRISTOL-MYERS SQUIBB COMPANY
LANGUAGE OF PUBL.:
                       English
DOCUMENT TYPE:
                       Patent
PATENT INFORMATION:
                       NUMBER
                                        KIND DATE
                       ______
                       WO 9902514
                                          A2 19990121
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                       CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE BF BJ
                       CF CG CI CM GA GN ML MR NE SN TD TG
PRIORITY INFO.:
                       US 1997-60/051,951
                                               19970708
                       US 1997-60/067,524
                                               19971204
APPLICATION INFO.:
                       WO 1998-US12550 A 19980616
      The present invention relates to compounds of formula (I), Q is selected
ABEN
       from the group
       consisting of (II), G is selected from the group consisting of alkyl,
       substituted alkyl, substituted
       or or unsubstituted aryl, heterocyclo, (III), W is O or NR15; X is O or
      H,H; Y is selected from the
       group consisting of O; H,OR16; OR17,OR17; NOR18; H,NOR19; H,NR20R21;
      H,H; or CHR22; OR17OR17 can be
       a cyclic ketal; Z1 and Z2 are selected from the group consisting of CH2,
      O, NR23, S or SO2, wherein
       only one of Z and Z2 is a heteroatom; B1 and B2 are selected from the
      group consisting of OR24, or
      OCOR25, or 2CNR26R27; when B1 is H and Y is OH, H they can form a
      six-membered ring ketal or acetal;
      D is selected from the group consisting of NR28R29, NR30COR31 or
       saturated heterocycle R1, R2, R3,
      R4, R5, R6, R7, R13, R14, R18, R19, R20, R21, R22, R26 and R27 are
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substituted alkyl, or aryl and when R1 and R2 are alkyl can be joined to form a cycloalkyl; R3 and R4 are alkyl can be joined to form a cycloalkyl; R9, R10, R16, R17, R24, R25, and R31 are selected from the group H, alkyl, or substituted alkyl; R8, R11, R12, R28, R30, R32, R33, and R30 are selected from the group consisting of H, alkyl, substituted alkyl, aryl, substituted aryl, cycloalkyl, or heterocyclo; R15, R23 and R29 are selected from the group consisting of H, alkyl, substituted alkyl, aryl, substituted aryl, cycloalkyl, heterocyclo, R32C=O, R33SO2, hydroxy, O-alkyl or O-substituted alkyl, the pharmaceutically acceptable salts thereof and any hydrates, solvates or geometric, optical and stereoisomers thereof, with the proviso that compounds wherein: W and X are both O; and R1, R2, R7 are H; and R3, R4, R6, are methyl; and R8, is H or methyl; and Z1, and Z2, are CH2; and G is 1-methyl-2-(substituted-4-thiazolyl)ethenyl; and Q is as defined above are excluded. La presente invention concerne des composes de la formule (I) dans laquelle Q est selectionne dans le groupe constitue par le groupement (II); G est selectionne dans le groupe constitue par alkyle, akyle substitue, aryle substitue ou insusbstitue, heterocyclo, le groupement (III); W est O ou NR15; X est O ou H,H; Y est selectionne dans le groupe constitue par O; H,OR16; OR17,OR17; NOR18; H, NOR19; H, NR20R21; H, H; ou CHR22; OR17, OR17 pouvant etre un cetal cyclique; Z1 et Z2 sont selectionnes dans le groupe constitue par CH2, O, NR23, S ou SO2, dans lequel seuls Z et Z2 sont un heteroatome; B1 et B2 sont selectionnes dans le groupe constitue par OR24 ou OCOR25 ou O2CNR26R27; et peuvent former ensemble un noyau cetal ou acetal a six chainons si B1 est H et Y est OH, H; D est selectionne dans le groupe constitue par NR28R29, NR30COR31 ou un heterocycle sature. R1, R2, R3, R4, R5, R6, R7, R13, R14, R18, R19, R20, R21, R22, R26 et R27 sont selectionnes dans le groupe constitue par H, alkyle, alkyle substitue ou aryle, et peuvent former ensemble un cycloalkyle si R1 et R2 ou R3 et R4 sont alkyle; R9, R10, R16, R17, R24, R25 et R31 sont selectionnes dans le groupe constitue par H, alkyle ou alkyle substitue; R8, R11, R12, R28, R30, R32, R33 et R30 sont selectionnes dans le groupe constitue par H, alkyle, alkyle substitue, aryle, aryle substitue, cycloalkyle ou heterocyclo; R15, R23 et R29 sont selectionnes dans le groupe constitue par H, alkyle, alkyle substitue, aryle, aryle substitue, cycloalkyle ou heterocyclo, R32C=O, R33SO2, hydroxy, O-alkyle ou O-alkyle substitue, leurs sels pharmaceutiquement acceptables ou leurs eventuels hydrates, solvates ou isomeres geometriques, optiques, ou stereoisomeres, a condition que soient exclus les composes dans lesquels W et X sont tous deux O; et R1, R2 et R7 sont H; et R3, R4 et R6 sont methyle; et R8 est H ou methyle; et Z1 et Z2 sont CH2; et G est

selected from the group H, alkyl,

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 $1-\text{methyl}-2-(\text{substitue}-4-\text{thiazolyl})\,\text{ethenyle;}$ et Q est tel que defini ci-dessus.

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=> d his
     (FILE 'HOME' ENTERED AT 10:09:50 ON 07 JUL 2008)
     FILE 'PCTFULL' ENTERED AT 10:10:27 ON 07 JUL 2008
           2484 S EPOTHILON?
L1
L2
           159 S L1/AB OR L1/TI
L3
             53 S L2 NOT PY>2001
L4
             33 S COMBINATION AND L3
L5
             29 S L4 AND (TAXOL OR PACLITAXEL)
             29 S L5 AND HER?
L6
L7
             1 S L5 AND (HER2 OR HER-2)
=> s 16 and (HER2 or HER-2)
         4722 HER2
        118696 HER
         1043 HERS
        119313 HER
                (HER OR HERS)
       1276185 2
          3260 HER-2
                (HER(W)2)
L8
             1 L6 AND (HER2 OR HER-2)
=> s 15 and (HER2 or HER-2)
         4722 HER2
        118696 HER
         1043 HERS
        119313 HER
                (HER OR HERS)
       1276185 2
          3260 HER-2
                (HER(W)2)
L9
             1 L5 AND (HER2 OR HER-2)
=> d ibib abs kwic
      ANSWER 1 OF 1
                       PCTFULL COPYRIGHT 2008 Univentio on STN
ACCESSION NUMBER:
                       1999002514 PCTFULL ED 20020515
TITLE (ENGLISH):
                       EPOTHILONE DERIVATIVES
TITLE (FRENCH):
                       DERIVES D'EPOTHILONE
                       VITE, Gregory, D.;
INVENTOR(S):
                        BORZILLERI, Robert, M.;
                        KIM, Soong-Hoon;
                        JOHNSON, James, A.
                       BRISTOL-MYERS SQUIBB COMPANY
PATENT ASSIGNEE(S):
LANGUAGE OF PUBL.:
                       English
DOCUMENT TYPE:
                        Patent
PATENT INFORMATION:
                       NUMBER
                                    KIND DATE
                        WO 9902514 A2 19990121
DESIGNATED STATES
       W:
                       AL AM AT AU AZ BA BB BG BR BY CA CH CN CU CZ DE DK EE
                        ES FI GB GE GH GM GW HU ID IL IS JP KE KG KP KR KZ LC
                        LK LR LS LT LU LV MD MG MK MN MW MX NO NZ PL PT RO RU
                        SD SE SG SI SK SL TJ TM TR TT UA UG UZ VN YU ZW GH GM
                        KE LS MW SD SZ UG ZW AM AZ BY KG KZ MD RU TJ TM AT BE
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PRIORITY INFO .:
                        US 1997-60/051,951
                                                19970708
                        US 1997-60/067,524
                                                19971204
                        WO 1998-US12550
APPLICATION INFO.:
                                             A 19980616
      The present invention relates to compounds of formula (I), Q is selected
ABEN
       from the group
       consisting of (II), G is selected from the group consisting of alkyl,
       substituted alkyl, substituted
       or or unsubstituted aryl, heterocyclo, (III), W is O or NR15; X is O or
       H,H; Y is selected from the
       group consisting of O; H,OR16; OR17,OR17; NOR18; H,NOR19; H,NR20R21;
       H,H; or CHR22; OR17OR17 can be
       a cyclic ketal; Z1 and Z2 are selected from the group consisting of CH2,
       O, NR23, S or SO2, wherein
       only one of Z and Z2 is a heteroatom; B1 and B2 are selected from the
       group consisting of OR24, or
       OCOR25, or 2CNR26R27; when B1 is H and Y is OH, H they can form a
       six-membered ring ketal or acetal;
       D is selected from the group consisting of NR28R29, NR30COR31 or
       saturated heterocycle R1, R2, R3,
       R4, R5, R6, R7, R13, R14, R18, R19, R20, R21, R22, R26 and R27 are
       selected from the group H, alkyl,
       substituted alkyl, or aryl and when R1 and R2 are alkyl can be joined to
       form a cycloalkyl; R3 and
       R4 are alkyl can be joined to form a cycloalkyl; R9, R10, R16, R17, R24,
       R25, and R31 are selected
       from the group H, alkyl, or substituted alkyl; R8, R11, R12, R28, R30,
       R32, R33, and R30 are
       selected from the group consisting of H, alkyl, substituted alkyl, aryl,
       substituted aryl,
       cycloalkyl, or heterocyclo; R15, R23 and R29 are selected from the
       group consisting of H, alkyl,
       substituted alkyl, aryl, substituted aryl, cycloalkyl, heterocyclo,
       R32C=O, R33SO2, hydroxy, O-alkyl
       or O-substituted alkyl, the pharmaceutically acceptable salts thereof
       and any hydrates, solvates or
       geometric, optical and stereoisomers thereof, with the proviso that
       compounds wherein: W and X are
       both O; and R1, R2, R7 are H; and R3, R4, R6, are methyl; and R8, is H
       or methyl; and Z1, and Z2,
       are CH2; and G is 1-methyl-2-(substituted-4-thiazolyl)ethenyl; and Q is
       as defined above are
       excluded.
       La presente invention concerne des composes de la formule (I) dans
ABFR
       laquelle Q est selectionne
       dans le groupe constitue par le groupement (II); G est selectionne dans
       le groupe constitue par
       alkyle, akyle substitue, aryle substitue ou insusbstitue, heterocyclo,
       le groupement (III); W est O
       ou NR15; X est O ou H,H; Y est selectionne dans le groupe constitue par
       O; H,OR16; OR17,OR17; NOR18;
       H, NOR19; H, NR20R21; H, H; ou CHR22; OR17, OR17 pouvant etre un cetal
       cyclique; Z1 et Z2 sont
       selectionnes dans le groupe constitue par CH2, O, NR23, S ou SO2, dans
       lequel seuls Z et Z2 sont un
       heteroatome; B1 et B2 sont selectionnes dans le groupe constitue par
       OR24 ou OCOR25 ou O2CNR26R27;
       et peuvent former ensemble un noyau cetal ou acetal a six chainons si B1
       est H et Y est OH, H; D est
       selectionne dans le groupe constitue par NR28R29, NR30COR31 ou un
       heterocycle sature. R1, R2, R3,
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CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE BF BJ

CF CG CI CM GA GN ML MR NE SN TD TG

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R4, R5, R6, R7, R13, R14, R18, R19, R20, R21, R22, R26 et R27 sont
       selectionnes dans le groupe
       constitue par H, alkyle, alkyle substitue ou aryle, et peuvent former
       ensemble un cycloalkyle si R1
       et R2 ou R3 et R4 sont alkyle; R9, R10, R16, R17, R24, R25 et R31 sont
       selectionnes dans le groupe
       constitue par H, alkyle ou alkyle substitue; R8, R11, R12, R28, R30,
       R32, R33 et R30 sont
       selectionnes dans le groupe constitue par H, alkyle, alkyle substitue,
       aryle, aryle substitue,
       cycloalkyle ou heterocyclo; R15, R23 et R29 sont selectionnes dans le
       groupe constitue par H,
       alkyle, alkyle substitue, aryle, aryle substitue, cycloalkyle ou
       heterocyclo, R32C=O, R33SO2,
       hydroxy, O-alkyle ou O-alkyle substitue, leurs sels pharmaceutiquement
       acceptables ou leurs
       eventuels hydrates, solvates ou isomeres geometriques, optiques, ou
       stereoisomeres, a condition que
       soient exclus les composes dans lesquels W et X sont tous deux O; et R1,
       R2 et R7 sont H; et R3, R4
       et R6 sont methyle; et R8 est H ou methyle; et Z1 et Z2 sont CH2; et G
       1-methyl-2-(substitue-4-thiazolyl)ethenyle; et Q est tel que defini
       ci-dessus.
       EPOTHILONE DERIVATIVES
TIEN
      DERIVES D'EPOTHILONE
TIFR
DETD
       R
       S Me
       ή0H
       N3 ], ]'] '
       0 Me
       0 OH 0
       I EpothiloneA R=H
       II EpothiloneB R=Me
       have been found to exert microtubule-stabilizing effects similar to
         TAXOL and hence cytotoxic activity against rapidly
       proliferating 'cells,
       such as, tumor cells or other hyperproliferative cellular disease, see
       .Angew. Chem. Int. Ed. Engl., . .
       The compounds of this invention. are also useful in combination
       with known anti-cancer and cytotoxic agents and treatments, including
       radiation. If formulated as a fixed dose, such combination
       products
       employ the compounds of this invention within the dosage range
       described below and the other pharmaceutically active agent within its
       approved dosage range. Compounds of formula V can be used
       sequentially with known anticancer or cytotoxic agents and treatment,
       including radiation when a combination formulation is
       inappropriate.
       Especially useful are cytotoxic drug combinations wherein the
       second
       drug chosen acts in a different phase of the cell cycle, e.g. S phase,
       than
       the present compounds of. . .
       Synthase Inhibitors,
       DNA Cross Linking Agents
       Topoisomerase I and II Inhibitors
```

DNA Alkylating Agents

Ribonucleoside Reductase Inhibitors
Cytotoxic Factors e.g. TNF-alpha or
Growth factor inhibitors e.g. HER 2 receptor MAB's
The present compounds may exist as multiple optical, geometric,
and stereoisomers. Included within the present invention are all such
isomers and. . .

• •

potency is accomplished following a modified procedure of Swindell, et al., (see Swindell, C.S., Krauss, N.E., Horwitz, S.B., and Ringel, I. Biologically active taxol analogues with deleted A-ring side chain substituents and

variable C-2' configurations. J. Med. Chem. 34: 1176-1184, 1991). These modifications, in part, result. . .

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cells were incubated at 37' form 72 hours at which time the tetrazolium dye, MTS at 333 $\rm gg/ml$ (final concentration), in combination

with the electron coupling agent phenazine methosulfate at 25 gm (final concentration) was added. A dehydrogenase enzyme in live cells reduces the MTS. . .

=>

---Logging off of STN---

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Executing the logoff script...

=> LOG Y

COST IN U.S. DOLLARS

FULL ESTIMATED COST

SINCE FILE TOTAL ENTRY SESSION 23.30 23.51

STN INTERNATIONAL LOGOFF AT 10:14:45 ON 07 JUL 2008